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 L3            20 L2

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L3 ANSWER 1 OF 20 CAPLUS COPYRIGHT 2006 ACS on STN  
 ACCESSION NUMBER: 2005:1315386 CAPLUS  
 DOCUMENT NUMBER: 144:45521  
 TITLE: Dual-acting serotonin-norepinephrine reuptake inhibitor (SNRI)-NMDA antagonists for the treatment of genitourinary disorders  
 INVENTOR(S): Thor, Karl Bruce  
 PATENT ASSIGNEE(S): Dynogen Pharmaceuticals, Inc., USA  
 SOURCE: PCT Int. Appl., 169 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2005117872	A2	20051215	WO 2005-US22897	20050603
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
US 2005282859	A1	20051222	US 2005-145022	20050603
PRIORITY APPLN. INFO.:			US 2004-576999P	P 20040604
			US 2004-607820P	P 20040907
			US 2004-640105P	P 20041228

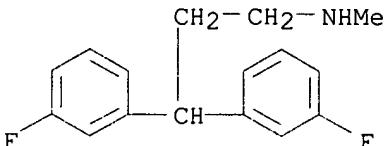
AB Compns. and methods are discloses for treatment of genitourinary disorders (e.g., urge incontinence). The compns. may generally include a dual-acting SNRI-NMDA antagonist (e.g., bicalafidine and/or milnacipran). Alternatively, the compns. may generally include an SNRI and an NMDA antagonist.

IT 186495-49-8 186495-55-6 186495-56-7

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)

RN 186495-49-8 CAPLUS

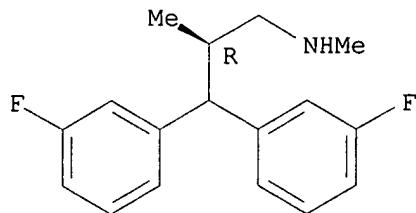
CN Benzenepropanamine, 3-fluoro- $\gamma$ -(3-fluorophenyl)-N-methyl- (9CI) (CA INDEX NAME)



RN 186495-55-6 CAPLUS

CN Benzenepropanamine, 3-fluoro- $\gamma$ -(3-fluorophenyl)-N, $\beta$ -dimethyl-,  
( $\beta$ R)- (9CI) (CA INDEX NAME)

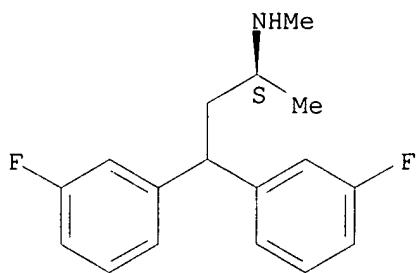
Absolute stereochemistry.



RN 186495-56-7 CAPLUS

CN Benzenepropanamine, 3-fluoro- $\gamma$ -(3-fluorophenyl)-N, $\alpha$ -dimethyl-,  
( $\alpha$ S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



IT Disease, animal

(Fowler's syndrome; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)

IT Glutamate antagonists

(NMDA antagonists; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)

IT Prostate gland, disease

(benign hyperplasia; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)

IT Hyperplasia

(benign prostatic; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)

IT Drug delivery systems

(buccal; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)

IT Drug delivery systems

(capsules; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)

IT Pain

(chronic pelvic pain syndrome; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)

IT Drug delivery systems

(controlled-release; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)

IT Bladder, disease

Inflammation

(cystitis, interstitial (cell); dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)

IT Drug delivery systems

(delayed release; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)

IT Urethra

(disease, urethritis; dual-acting serotonin-norepinephrine reuptake

IT inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
5-HT reuptake inhibitors  
Analgesics  
Anti-inflammatory agents  
Antitumor agents  
Bladder, disease  
Combination chemotherapy  
Drug delivery systems  
Human  
Neoplasm  
Urogenital system, disease  
(dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
IT Bladder, disease  
(hyperreflexia; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
IT Bladder, disease  
(incontinence; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
IT Drug delivery systems  
(inhaled; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
IT Drug delivery systems  
(intravesical; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
IT Drug delivery systems  
(mucosal; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
IT Nervous system agents  
(noradrenaline reuptake inhibitors; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
IT Drug delivery systems  
(oral; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
IT Testis  
(orchidalgia; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
IT Bladder, disease  
(overactive bladder, including overactive bladder with sphincter dysfunction; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
IT Disease, animal  
(pelvic hypersensitivity or sphincteric spasticity; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
IT Prostate gland, disease  
(prostodynia; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
IT Inflammation  
Prostate gland, disease  
(prostatitis; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
IT Drug delivery systems  
(pulsatile-release; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
IT Drug delivery systems  
(rectal; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
IT Bladder  
(smooth muscle; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
IT Muscle  
(smooth, urinary bladder; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
IT Drug delivery systems  
(sublingual; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
IT Drug delivery systems

(sustained-release; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
 IT Drug delivery systems  
 (tablets; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
 IT Drug delivery systems  
 (transdermal; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
 IT Drug delivery systems  
 (transurethral; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
 IT Urethra  
 (urethra stricture disease; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
 IT Inflammation  
 (urethritis; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
 IT Reproductive system  
 (vulva, vulvodynia or vulvar vestibulitis; dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA antagonists for treatment of genitourinary disorders)  
 IT 5586-73-2 14451-09-3, 5H-Dibenzo[a,d]cycloheptene-5-ethanamine  
 21745-77-7, 9H-Xanthene-9-ethanamine 21745-81-3, 9H-Thioxanthene-9-  
 ethanamine 21745-82-4 21745-85-7 28075-29-8 57226-64-9  
 63106-93-4 63940-51-2 66504-40-3 69096-48-6 71195-57-8  
 92623-85-3 105310-27-8 109306-10-7 136090-96-5 136090-97-6  
 144451-98-9 153275-06-0 170018-66-3 170018-67-4 170018-79-8  
 170018-83-4 186495-47-6 186495-48-7 **186495-49-8**  
 186495-52-3 186495-53-4 186495-54-5 **186495-55-6**  
**186495-56-7** 186495-66-9 186495-67-0 186495-80-7  
 186495-84-1 186495-86-3 186495-87-4 186495-88-5 186495-89-6  
 186495-90-9 186496-20-8 186496-23-1 186496-29-7 186496-30-0  
 186496-71-9 200429-73-8 200429-74-9 200429-75-0 200429-79-4  
 200429-80-7 200429-81-8 200429-82-9 200429-83-0 200429-84-1  
 200429-85-2 200429-86-3 200429-87-4 255039-66-8 255039-67-9  
 255039-68-0 255039-69-1 255039-71-5 255039-73-7 255039-75-9  
 255039-77-1 255039-79-3 255039-81-7 255040-00-7 255040-01-8  
 255040-02-9 255040-03-0 255040-04-1 255040-05-2 255040-07-4  
 255040-08-5 410074-73-6 410074-75-8 435293-68-8 688738-11-6  
 688738-12-7 871100-17-3 871100-18-4 871100-19-5 871100-20-8  
 871100-21-9 871100-22-0 871100-23-1 871331-21-4 871331-22-5  
 871331-23-6 871331-24-7 871331-25-8  
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
 (Biological study); USES (Uses)  
 (dual-acting serotonin-norepinephrine reuptake inhibitor-NMDA  
 antagonists for treatment of genitourinary disorders)

L3 ANSWER 2 OF 20 CAPLUS COPYRIGHT 2006 ACS on STN  
 ACCESSION NUMBER: 2005:409336 CAPLUS  
 DOCUMENT NUMBER: 142:457117  
 TITLE: Neuroprotective effects of gly-pro-glu following  
 intravenous infusion  
 INVENTOR(S): Guan, Jian; Thomas, Gregory Brian; Batchelor, David  
 Charles; Gluckman, Peter David  
 PATENT ASSIGNEE(S): Neuren Pharmaceuticals Ltd., N. Z.; Neuren  
 Pharmaceuticals Inc.  
 SOURCE: PCT Int. Appl., 48 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2005042000	A1	20050512	WO 2004-US35165	20041022
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,				

GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,  
 LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,  
 NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,  
 TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW  
 RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,  
 AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,  
 EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE,  
 SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE,  
 SN, TD, TG

PRIORITY APPLN. INFO.: US 2003-513851P P 20031023  
 US 2003-515397P P 20031028  
 US 2004-553688P P 20040316

**AB** Gly-Pro-Glu (GPE) is rapidly metabolized in vivo. We found that GPE infusion elicits potent and consistent neuroprotection in all brain regions examined, and in certain embodiments, the effects were greater than those of a bolus injection followed by infusion ('loading dose/infusion'). GPE reduced apoptosis in the hippocampus and inhibited microglial proliferation and prevented the injury-induced loss of astrocytes and improved long-term somatofunction. GPE after infusion showed a broad ED range (0.3-30mg/kg/h) and had a surprisingly extended window of treatment efficacy, permitting its use from 1 to at least as late as 24 h after neural injury. We also found that neuroprotective effects of acute GPE administration were prolonged and therefore capable of being used effectively to treat a variety of neurodegenerative conditions, even when administered after a neural injury. Thus, GPE can be an effective neuroprotective agent used either alone or co-administered along with other neuroprotective agents, antiinflammatory agents or peptidase or protease inhibitors. Compns. of GPE and protease and/or peptidase inhibitors are provided.

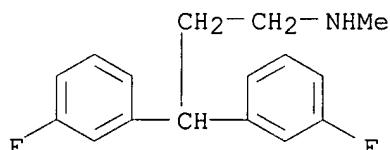
**IT** 186495-99-8, NPS 1506

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(neuroprotective effects of gly-pro-glu following i.v. infusion)

186495-99-8 CAPLUS

Benzene propanamine, 3-fluoro- $\gamma$ -(3-fluorophenyl)-N-methyl-, hydrochloride (9CI) (CA INDEX NAME)



● HCl

**IT** Bone morphogenetic proteins

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(2; neuroprotective effects of gly-pro-glu following i.v. infusion)

**IT** AIDS (disease)

(AIDS dementia complex; neuroprotective effects of gly-pro-glu following i.v. infusion)

**IT** Mental and behavioral disorders

(AIDS dementia; neuroprotective effects of gly-pro-glu following i.v. infusion)

**IT** Brain, disease

Prion diseases

(Creutzfeldt-Jakob; neuroprotective effects of gly-pro-glu following i.v. infusion)

**IT** Growth factors, animal

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(Glial activating factor; neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Gene, animal  
RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(HSTF1; neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Nervous system, disease  
(Huntington's chorea; neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Insulin-like growth factor-binding proteins  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(IGFBP-3; neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Integrins  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(LPAM-1 (lymphocyte Peyer's patch adhesion mol. 1); neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Antibodies and Immunoglobulins  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(MAdCAM-1; neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Brain, disease  
(Schilder's disease; neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Nervous system, disease  
(amyotrophic lateral sclerosis; neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Brain  
(cerebral cortex; neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Ischemia  
(cerebral; neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Encephalomyelitis  
(chronic relapsing; neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Surgery  
(coronary artery bypass; neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Artery  
(coronary, bypass surgery; neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Brain  
(corpus striatum; neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Radiation  
(damage; neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Nerve, disease  
(death; neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Nerve, disease  
Nervous system, disease  
(degeneration; neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Central nervous system, disease  
(demyelination; neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Brain  
(dentate gyrus; neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Mental and behavioral disorders  
(depression; neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Surgery  
(elective; neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Neurotrophic factors  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(glial-derived; neuroprotective effects of gly-pro-glu following i.v.  
infusion)

IT Injury  
(head and neck; neuroprotective effects of gly-pro-glu following i.v.  
infusion)

IT Brain  
(hippocampus, sector CA1; neuroprotective effects of gly-pro-glu  
following i.v. infusion)

IT Brain  
(hippocampus, sector CA2; neuroprotective effects of gly-pro-glu  
following i.v. infusion)

IT Brain  
(hippocampus, sector CA3; neuroprotective effects of gly-pro-glu  
following i.v. infusion)

IT Brain  
(hippocampus, sector CA4; neuroprotective effects of gly-pro-glu  
following i.v. infusion)

IT Drug delivery systems  
(infusions, i.v.; neuroprotective effects of gly-pro-glu following i.v.  
infusion)

IT Head and Neck, disease  
Nerve, disease  
Reperfusion  
(injury; neuroprotective effects of gly-pro-glu following i.v.  
infusion)

IT Gene, animal  
RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL  
(Biological study); USES (Uses)  
(int-2; neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Brain, disease  
Nerve, disease  
(ischemia; neuroprotective effects of gly-pro-glu following i.v.  
infusion)

IT Brain, disease  
(leukoencephalopathy; neuroprotective effects of gly-pro-glu following  
i.v. infusion)

IT Neuroglia  
(microglia; neuroprotective effects of gly-pro-glu following i.v.  
infusion)

IT Inflammation  
Spinal cord, disease  
(myelitis; neuroprotective effects of gly-pro-glu following i.v.  
infusion)

IT Inflammation  
Nerve, disease  
(neuritis; neuroprotective effects of gly-pro-glu following i.v.  
infusion)

IT Nerve, neoplasm  
(neuroblastoma; neuroprotective effects of gly-pro-glu following i.v.  
infusion)

IT Cell death  
(neuron; neuroprotective effects of gly-pro-glu following i.v.  
infusion)

IT Injury  
Ischemia  
(neuronal; neuroprotective effects of gly-pro-glu following i.v.  
infusion)

IT Alzheimer's disease  
Anti-Alzheimer's agents  
Anti-inflammatory agents  
Anticonvulsants  
Antidepressants  
Antiparkinsonian agents  
Antipsychotics  
Asphyxia  
Astrocyte  
Down's syndrome  
Encephalitis  
Encephalomyelitis

Epilepsy  
Hypoxia  
Inflammation  
Leukemia  
Meningitis  
Multiple sclerosis  
Parkinson's disease  
Schizophrenia  
Spinal muscular atrophy  
(neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Toxins  
RL: ADV (Adverse effect, including toxicity); BIOL (Biological study)  
(neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Cytokines  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Proliferating cell nuclear antigen  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Insulin-like growth factor-binding proteins

Interleukins  
Neurotrophic factors  
Tumor necrosis factors  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
(Biological study); USES (Uses)  
(neuroprotective effects of gly-pro-glu following i.v. infusion)

IT Cytoprotective agents  
(neuroprotective; neuroprotective effects of gly-pro-glu following i.v.  
infusion)

IT Nervous system, disease  
(optic neuritis; neuroprotective effects of gly-pro-glu following  
i.v. infusion)

IT Nerve, disease  
(peripheral neuropathy; neuroprotective effects of gly-pro-glu  
following i.v. infusion)

IT Brain, disease  
(progressive multifocal leukoencephalopathy; neuroprotective effects of  
gly-pro-glu following i.v. infusion)

IT Paralysis  
(pseudobulbar; neuroprotective effects of gly-pro-glu following i.v.  
infusion)

IT Injury  
(reperfusion; neuroprotective effects of gly-pro-glu following i.v.  
infusion)

IT Proteins  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
(Biological study); USES (Uses)  
(somatotropin-binding; neuroprotective effects of gly-pro-glu following  
i.v. infusion)

IT Brain, disease  
Brain, disease  
(stroke; neuroprotective effects of gly-pro-glu following i.v.  
infusion)

IT Brain, disease  
(trauma; neuroprotective effects of gly-pro-glu following i.v.  
infusion)

IT Interferons  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
(Biological study); USES (Uses)  
( $\alpha$ ; neuroprotective effects of gly-pro-glu following i.v.  
infusion)

IT Integrins  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
( $\alpha 4\beta 1$ ; neuroprotective effects of gly-pro-glu following i.v.  
infusion)

IT Transforming growth factors  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
(Biological study); USES (Uses)  
( $\beta 1$ ; neuroprotective effects of gly-pro-glu following i.v.

IT infusion)  
IT Interferons  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
( $\beta$ ; neuroprotective effects of gly-pro-glu following i.v.  
infusion)  
IT Interferons  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
( $\gamma$ ; neuroprotective effects of gly-pro-glu following i.v.  
infusion)  
IT 9001-92-7, Proteinase  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(inhibitors; neuroprotective effects of gly-pro-glu following i.v.  
infusion)  
IT 9015-82-1, Peptidyldipeptidase 9031-94-1, Aminopeptidase 9031-98-5,  
Carboxypeptidase 9031-99-6, Dipeptidase  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(inhibitor; neuroprotective effects of gly-pro-glu following i.v.  
infusion)  
IT 56-40-6, Glycine, biological studies 56-86-0, L-Glutamic acid,  
biological studies 147-85-3, Proline, biological studies 704-15-4  
RL: ANT (Analyte); PKT (Pharmacokinetics); ANST (Analytical study); BIOL (Biological study)  
(neuroprotective effects of gly-pro-glu following i.v. infusion)  
IT 37205-61-1, Proteinase inhibitor 37259-58-8, Serine protease  
37353-41-6, Cysteine protease 123584-45-2, Fibroblast growth factor 4  
130939-41-2, Fibroblast growth factor 6 145266-99-5, Metalloproteinase  
inhibitor 148348-14-5, Fibroblast growth factor 3 169592-56-7,  
Caspase-3  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(neuroprotective effects of gly-pro-glu following i.v. infusion)  
IT 32302-76-4  
RL: PAC (Pharmacological activity); PKT (Pharmacokinetics); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(neuroprotective effects of gly-pro-glu following i.v. infusion)  
IT 492-27-3, Kynurenic acid 533-45-9, Clomethiazole 9002-72-6, Growth  
hormone 9061-61-4, Nerve growth factor 9087-70-1, Aprotinin  
14611-51-9, Selegiline 26305-03-3, Pepstatin A 30827-99-7, AEBSF  
50913-82-1, ORG 2766 55123-66-5, Leupeptin 58970-76-6, Bestatin  
66701-25-5 67763-96-6, IGF-1 67763-97-7, IGF-II 77086-22-7, MK-801  
80714-61-0, Semax 104987-11-3, Tacrolimus 106096-92-8, Acidic  
fibroblast growth factor 106096-93-9, Basic fibroblast growth factor  
106956-32-5, Oncostatin M 114949-22-3, Activin 118876-58-7, NBQX  
130939-66-1, Neurotrophin 3 143375-33-1, Neurotrophin 4 148348-15-6,  
Keratinocyte growth factors 161832-65-1, LY 300164 161832-71-9, LY  
303070 171758-70-6, Keratinocyte growth factor 2 186495-99-8,  
NPS 1506 204719-95-9, Fibroblast growth factor 16 524706-48-7, GV  
1505260  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(neuroprotective effects of gly-pro-glu following i.v. infusion)  
REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 3 OF 20 CAPLUS COPYRIGHT 2006 ACS on STN  
ACCESSION NUMBER: 2004:1059129 CAPLUS  
DOCUMENT NUMBER: 142:32998  
TITLE: Compositions of a cyclooxygenase-2 selective inhibitor  
and a cannabinoid agent for the treatment of central  
nervous system damage  
INVENTOR(S): Stephenson, Diane T.; Taylor, Duncan P.  
PATENT ASSIGNEE(S): Pharmacia Corporation, USA  
SOURCE: PCT Int. Appl., 177 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004105699	A2	20041209	WO 2004-US16496	20040526
WO 2004105699	A3	20051215		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				

PRIORITY APPLN. INFO.: US 2003-473820P P 20030528

OTHER SOURCE(S): MARPAT 142:32998

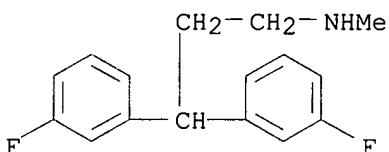
AB The present invention provides compns. and methods for the treatment of central nervous system damage in a subject. More particularly, the invention provides a combination therapy for the treatment of a central nervous system ischemic condition or a central nervous system traumatic injury comprising the administration to a subject of a cannabinoid agent in combination with a cyclooxygenase-2 selective inhibitor.

IT 186495-49-8, Delucemine

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(compns. of a cyclooxygenase-2 selective inhibitor and a cannabinoid agent for treatment of central nervous system damage)

RN 186495-49-8 CAPLUS

CN Benzenepropanamine, 3-fluoro- $\gamma$ -(3-fluorophenyl)-N-methyl- (9CI) (CA INDEX NAME)



IT Ischemia

(central nervous system; compns. of a selective cyclooxygenase-2 selective inhibitor and a cannabinoid agent for treatment of central nervous system damage)

IT Combination chemotherapy

Drug interactions

Ischemia

(compns. of a selective cyclooxygenase-2 selective inhibitor and a cannabinoid agent for treatment of central nervous system damage)

IT Cannabinoids

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(compns. of a selective cyclooxygenase-2 selective inhibitor and a cannabinoid agent for treatment of central nervous system damage)

IT Central nervous system, disease

(ischemia; compns. of a selective cyclooxygenase-2 selective inhibitor and a cannabinoid agent for treatment of central nervous system damage)

IT Cytoprotective agents

(neuroprotective; compns. of a selective cyclooxygenase-2 selective inhibitor and a cannabinoid agent for treatment of central nervous system damage)

IT Brain, disease

(stroke; compns. of a selective cyclooxygenase-2 selective inhibitor and a cannabinoid agent for treatment of central nervous system damage)

IT 52-52-8, 1-Aminocyclopentanecarboxylic acid 56-40-6, Glycine, biological studies 83-98-7, Orphenadrine 521-35-7, Cannabinol 726-99-8, Fluorofelbamate 1972-08-3, Dronabinol 7541-16-4 13956-29-1,

Cannabidiol 25451-15-4, Felbamate 35377-89-0, 1-Methoxy-endo-4-hydroxy-9-oxabicyclo[3.3.1]nonane 38964-50-0 53847-30-6, 2-Arachidonylglycerol 57982-78-2, Budipine 68134-81-6, Gacyclidine 71125-38-7, Meloxicam 76163-84-3 76163-85-4 76163-87-6 76163-88-7 80286-75-5 83002-04-4 92623-85-3, Milnacipran 93438-65-4, Conantokin G 96206-92-7, 2-Methyl-6-(phenylethynyl)pyridine 97240-79-4, Topiramate 104454-71-9, Ipenoxazone 112924-45-5, Dexanabinol 117414-74-1, Midafotel 117571-54-7 119784-07-5 120667-19-8 123653-11-2, N-[2-(Cyclohexyloxy)-4-nitrophenyl]methanesulfonamide 124649-81-6 128298-28-2, Remacemide 132472-31-2 135025-56-8, 7-Chlorothiokynurenic acid 136109-04-1 137159-92-3, Aptiganel 138047-56-0 139051-78-8 140835-14-9 142235-88-9 143850-75-3 144912-63-0 153322-05-5, Lanicemine 153504-81-5, Licostinel 155471-08-2 157182-49-5 158328-22-4 160754-76-7 161230-88-2 161292-39-3 162011-90-7, Rofecoxib 164178-33-0 166974-22-7 168273-06-1 169590-41-4, Deracoxib 169590-42-5, Celecoxib 173186-99-7 176977-56-3, [6-Methoxy-2-(4-methoxyphenyl)benzo[b]furan-3-yl](4-cyanophenyl)methanone 180200-68-4, 4-(4-Cyclohexyl-2-methyloxazol-5-yl)-2-fluorobenzenesulfonamide 181695-72-7, Valdecoxib 183232-66-8 186495-49-8, Delucemine 192703-06-3 193278-48-7 193356-17-1 197077-52-4 197438-41-8 198470-84-7, Parecoxib 198710-92-8, Kaitocephalin 200430-63-3 202409-33-4, Etoricoxib 202463-68-1 202807-80-5 202914-18-9 212126-32-4, 2-(3,5-Difluorophenyl)-3-[4-(methylsulfonyl)phenyl]-2-cyclopenten-1-one 215123-80-1 219810-59-0, Neramexane 220991-20-8, Lumiracoxib 220991-33-3 252374-41-7 253450-09-8, Besonprodil 256510-26-6 266320-83-6 342047-49-8 369640-27-7 803731-69-3 803731-70-6 803731-71-7  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(compns. of a cyclooxygenase-2 selective inhibitor and a cannabinoid agent for treatment of central nervous system damage)  
IT 329900-75-6, Cyclooxygenase 2 329967-85-3, Cyclooxygenase-1  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(compns. of a selective cyclooxygenase-2 selective inhibitor and a cannabinoid agent for treatment of central nervous system damage)

L3 ANSWER 4 OF 20 CAPLUS COPYRIGHT 2006 ACS on STN  
ACCESSION NUMBER: 2004:781548 CAPLUS  
DOCUMENT NUMBER: 141:254405  
TITLE: Acute treatment with MgSO<sub>4</sub> attenuates long-term hippocampal tissue loss after brain trauma in the rat  
AUTHOR(S): Browne, Kevin D.; Leoni, Matthew J.; Iwata, Akira; Chen, Xiao-Han; Smith, Douglas H.  
CORPORATE SOURCE: Department of Neurosurgery, University of Pennsylvania, Philadelphia, PA, USA  
SOURCE: Journal of Neuroscience Research (2004), 77(6), 878-883  
CODEN: JNREDK; ISSN: 0360-4012  
PUBLISHER: Wiley-Liss, Inc.  
DOCUMENT TYPE: Journal  
LANGUAGE: English  
AB Previous studies have shown that magnesium salts and the noncompetitive N-methyl-D-aspartate (NMDA) receptor antagonist, NPS 1506, attenuated short-term cognitive deficits and histopathol. changes associated with traumatic brain injury (TBI). We evaluated the long-term effects of both therapies after brain trauma. Young adult rats were subjected to parasagittal fluid-percussion brain injury and received either MgSO<sub>4</sub> (125 μmol/400 g rat; n = 12) 15 min post-injury, NPS 1506 (1.15 mg/kg; n = 12) 15 min and 4 h post-injury, or vehicle (n = 9) 15 min post-injury. Uninjured animals (sham) received vehicle (n = 10). Learning function in these animals was evaluated using a water maze paradigm 8 mo after injury or sham treatment, and the brains were examined for cortical and hippocampal tissue loss. Compared to sham animals, injured vehicle-treated animals displayed a substantial learning dysfunction, indicated by an increased latency to find a hidden platform in the water maze (P < 0.001). No improvements in learning, however, were found for injured animals treated with NPS 1506 or MgSO<sub>4</sub>. Injury induced >30% loss of tissue in the ipsilateral cortex in vehicle-treated animals that was not reduced in animals treated with either NPS 1506 or MgSO<sub>4</sub>. Treatment with MgSO<sub>4</sub>

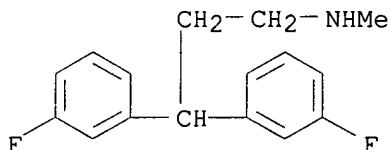
significantly reduced progressive tissue loss in the hippocampus ( $P < 0.001$ ). These findings are the first to demonstrate long-term neuroprotection of hippocampal tissue by an acute treatment in a TBI model. These data also show that the previously reported broad efficacy of MgSO<sub>4</sub> or NPS 1506 observed shortly after brain trauma could not be detected 8 mo post-injury.

IT 186495-99-8, NPS 1506

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(MgSO<sub>4</sub> attenuates long-term hippocampal tissue loss after brain trauma in rat)

RN 186495-99-8 CAPLUS

CN Benzenepropanamine, 3-fluoro- $\gamma$ -(3-fluorophenyl)-N-methyl-, hydrochloride (9CI) (CA INDEX NAME)



● HCl

IT Cognition

Learning disorders

(MgSO<sub>4</sub> attenuates long-term hippocampal tissue loss after brain trauma in rat)

IT Glutamate antagonists

(NMDA antagonists; MgSO<sub>4</sub> attenuates long-term hippocampal tissue loss after brain trauma in rat)

IT Brain

(cortex; MgSO<sub>4</sub> attenuates long-term hippocampal tissue loss after brain trauma in rat)

IT Brain

(hippocampus, atrophy; MgSO<sub>4</sub> attenuates long-term hippocampal tissue loss after brain trauma in rat)

IT Cytoprotective agents

(neuroprotective; MgSO<sub>4</sub> attenuates long-term hippocampal tissue loss after brain trauma in rat)

IT Brain, disease

(trauma; MgSO<sub>4</sub> attenuates long-term hippocampal tissue loss after brain trauma in rat)

IT 7487-88-9, Sulfuric acid magnesium salt (1:1), biological studies

186495-99-8, NPS 1506

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(MgSO<sub>4</sub> attenuates long-term hippocampal tissue loss after brain trauma in rat)

REFERENCE COUNT: 43 THERE ARE 43 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 5 OF 20 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:353140 CAPLUS

DOCUMENT NUMBER: 140:380634

TITLE: Compositions of cyclooxygenase-2 selective inhibitors and NMDA receptor antagonist for the treatment or prevention of neuropathic pain

INVENTOR(S): Cheung, Raymond Y.

PATENT ASSIGNEE(S): Pharmacia Corporation, USA

SOURCE: U.S. Pat. Appl. Publ., 51 pp.

CODEN: USXXCO

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2004082543	A1	20040429	US 2002-282660	20021029
WO 2004039371	A2	20040513	WO 2003-US33089	20031017
WO 2004039371	A3	20040617		
			W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG	

PRIORITY APPLN. INFO.: US 2002-282660 A 20021029

OTHER SOURCE(S): MARPAT 140:380634

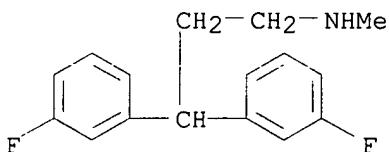
AB The present invention provides compns. and methods to treat or prevent neuropathic pain in a subject using a combination of a COX-2 selective inhibitor and a NMDA receptor antagonist.

IT 186495-49-8, Delucemine

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (compns. of cyclooxygenase-2 selective inhibitors and NMDA receptor antagonists for treatment or prevention of neuropathic pain)

RN 186495-49-8 CAPLUS

CN Benzenepropanamine, 3-fluoro-γ-(3-fluorophenyl)-N-methyl- (9CI) (CA INDEX NAME)



IT Glutamate receptors

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (NMDA-binding; compns. of cyclooxygenase-2 selective inhibitors and NMDA receptor antagonists for treatment or prevention of neuropathic pain)

IT Pain

(neuropathic; compns. of cyclooxygenase-2 selective inhibitors and NMDA receptor antagonists for treatment or prevention of neuropathic pain)

IT Nerve, disease

(neuropathy, related pain, treatment of; compns. of cyclooxygenase-2 selective inhibitors and NMDA receptor antagonists for treatment or prevention of neuropathic pain)

IT Drug delivery systems

(prodrugs; compns. of cyclooxygenase-2 selective inhibitors and NMDA receptor antagonists for treatment or prevention of neuropathic pain)

IT 329900-75-6, COX-2

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (COX-2; compns. of cyclooxygenase-2 selective inhibitors and NMDA receptor antagonists for treatment or prevention of neuropathic pain)

IT 52-52-8, 1-Aminocyclopentane-carboxylic acid 56-40-6, Glycine, biological studies 83-98-7, Orphenadrine 125-71-3, Dextromethorphan 254-04-6, 2H-1-Benzopyran 726-99-8, Fluorofelbamate 768-94-5, Amantadine 6740-88-1, Ketamine 19982-08-2, Memantine 25451-15-4, Felbamate 57982-78-2, Budipine 68134-81-6, Gacyclidine 70172-33-7 71125-38-7, Meloxicam 92623-85-3, Milnacipran 93438-65-4, Conantokin G 96206-92-7, 2-Methyl-6-(phenylethynyl)-pyridine 97240-79-4, Topiramate 104454-71-9, Ipenoxazone 112924-45-5, Dexanabinol 117414-74-1, Midafotel 117571-54-7 120667-19-8 123653-11-2, {N-[2-(Cyclohexyloxy)-4-nitrophenyl]methanesulfonamide} 128298-28-2, Remacemide 132472-31-2 134234-12-1, Traxoprodil 135025-56-8, 7-Chlorothiokynurenic acid

136109-04-1 137159-92-3, Aptiganel 138047-56-0 139051-78-8  
 142235-88-9 143850-75-3 144912-63-0 153322-05-5, Lanicemine  
 153504-81-5, Licostinel 160754-76-7 161230-88-2 161292-39-3  
 162011-90-7, Rofecoxib 166974-22-7 169590-41-4, Deracoxib  
 169590-42-5, Celecoxib 170029-85-3 173186-99-7 180200-68-4,  
 4-(4-Cyclohexyl-2-methyloxazol-5-yl)-2-fluorobenzenesulfonamide  
 181695-72-7, Valdecoxib 186495-49-8, Delucemine 193278-48-7  
 193356-17-1 197077-52-4 198470-84-7, Parecoxib 198710-92-8,  
 Kaitocephalin 200430-63-3 202409-33-4, Etoricoxib 202807-80-5  
 202914-18-9 212126-32-4, 2-(3,5-Difluorophenyl)-3-[4-  
 (methylsulfonyl)phenyl]-2-cyclopenten-1-one 219810-59-0, Neramexane  
 252374-41-7 253450-09-8, Besonprodil 266320-83-6, ABT 963  
 342047-49-8 369640-27-7 676451-52-8D, derivs.  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (compns. of cyclooxygenase-2 selective inhibitors and NMDA receptor  
 antagonists for treatment or prevention of neuropathic pain)

L3 ANSWER 6 OF 20 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2003:924889 CAPLUS

DOCUMENT NUMBER:

140:317215

TITLE: Synthesis and brain regional distribution of [11C]NPS  
1506 in mice and rat: An N-methyl-D-aspartate (NMDA)  
receptor antagonist

AUTHOR(S): Fuchigami, Takeshi; Haradahira, Terushi; Arai, Takuya;  
Okauchi, Takashi; Maeda, Jun; Suzuki, Kazutoshi;  
Yamamoto, Fumihiro; Suhara, Tetsuya; Sasaki, Shigeki;  
Maeda, Minoru

CORPORATE SOURCE: Graduate School of Pharmaceutical Sciences, Kyushu  
University, Fukuoka, 812-8582, Japan

SOURCE: Biological & Pharmaceutical Bulletin (2003), 26(11),  
1570-1573

PUBLISHER: CODEN: BPBLEO; ISSN: 0918-6158  
Pharmaceutical Society of Japan

DOCUMENT TYPE: Journal

LANGUAGE: English

AB NPS 1506 [3-fluoro- $\gamma$ -(3-fluorophenyl)-N-methylbenzenepropamine] is representative of a non-psychotomimetic class of N-methyl-D-aspartate (NMDA) receptor antagonists. [11C]NPS 1506 was prepared at high radiochem. purity (>98%) with a specific activity of around 50 GBq/ $\mu$ mol at the end of synthesis by methylation of the desmethyl precursor with [11C]methyl iodide in the presence of NaH. Biodistribution of [11C]NPS 1506 in mice and rat demonstrated that uptake into the brain was rapid and occurred at high levels. [11C]NPS 1506 showed no appreciable specific binding in rodent brains under in vivo conditions, possibly because of both a large non-specific bound fraction and low in vitro binding affinity for NMDA receptors.

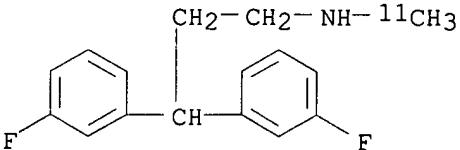
IT 677764-00-0P

RL: PKT (Pharmacokinetics); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)

(NMDA receptor antagonist [11C]NPS 1506 preparation and brain distribution  
in mice and rat)

RN 677764-00-0 CAPLUS

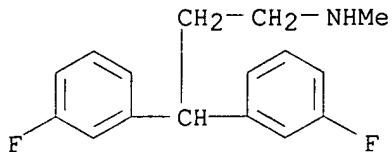
CN Benzene propanamine, 3-fluoro- $\gamma$ -(3-fluorophenyl)-N-(methyl-11C)-, hydrochloride (9CI) (CA INDEX NAME)



● HCl

IT 186495-99-8P, NPS 1506

RL: PRP (Properties); RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)  
(NMDA receptor antagonist [11C]NPS 1506 preparation and brain distribution  
in mice and rat)  
RN 186495-99-8 CAPLUS  
CN Benzenepropanamine, 3-fluoro- $\gamma$ -(3-fluorophenyl)-N-methyl-,  
hydrochloride (9CI) (CA INDEX NAME)



● HCl

IT Glutamate antagonists  
(NMDA antagonists; NMDA receptor antagonist [11C]NPS 1506 preparation and  
brain distribution in mice and rat)  
IT Blood-brain barrier  
Brain  
Isotope indicators  
(NMDA receptor antagonist [11C]NPS 1506 preparation and brain distribution  
in mice and rat)  
IT Glutamate receptors  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(NMDA-binding; NMDA receptor antagonist [11C]NPS 1506 preparation and brain  
distribution in mice and rat)  
IT 677764-00-0P  
RL: PKT (Pharmacokinetics); SPN (Synthetic preparation); BIOL (Biological  
study); PREP (Preparation)  
(NMDA receptor antagonist [11C]NPS 1506 preparation and brain distribution  
in mice and rat)  
IT 186495-99-8P, NPS 1506  
RL: PRP (Properties); RCT (Reactant); SPN (Synthetic preparation); PREP  
(Preparation); RACT (Reactant or reagent)  
(NMDA receptor antagonist [11C]NPS 1506 preparation and brain distribution  
in mice and rat)  
IT 542-92-7, Cyclopentadiene, reactions 170019-10-0  
RL: RCT (Reactant); RACT (Reactant or reagent)  
(NMDA receptor antagonist [11C]NPS 1506 preparation and brain distribution  
in mice and rat)  
IT 677763-98-3P  
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT  
(Reactant or reagent)  
(NMDA receptor antagonist [11C]NPS 1506 preparation and brain distribution  
in mice and rat)

REFERENCE COUNT: 16 THERE ARE 16 CITED REFERENCES AVAILABLE FOR THIS  
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 7 OF 20 CAPLUS COPYRIGHT 2006 ACS on STN  
ACCESSION NUMBER: 2003:396671 CAPLUS  
DOCUMENT NUMBER: 138:379256  
TITLE: Cyclic prolylglycine composition and therapeutic uses  
INVENTOR(S): Tran, Loi  
PATENT ASSIGNEE(S): USA  
SOURCE: PCT Int. Appl., 37 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 2  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
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WO 2003041655	A2	20030522	WO 2002-US36639	20021112
WO 2003041655	A3	20040910		
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW			
RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG			

CA 2466701	AA	20030522	CA 2002-2466701	20021112
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US 2003109531	A1	20030612	US 2002-292732	20021112
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PRIORITY APPLN. INFO.:

NZ 2001-515432	A	20011113
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US 2002-405909P	P	20020826
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WO 2002-US36639	W	20021112
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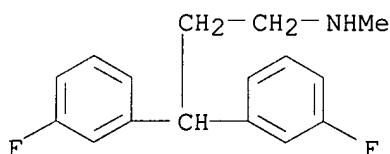
AB The invention discloses compns. containing, and use of, cyclic prolylglycine, and analogs and mimetics thereof, as neuroprotective agents for the treatment and or prevention of neurol. disorders including but not limited to cerebral ischemia or cerebral infarction resulting from a range of phenomena, e.g. thromboembolic or hemorrhagic stroke, cerebral basospasms, hypoglycemia, cardiac arrest, status epilepticus, perinatal asphyxia, anoxia (e.g. from drowning), pulmonary surgery, and cerebral trauma, as well as the treatment and prevention of chronic neurodegenerative disorders, e.g. Alzheimer's disease, Parkinson's disease, and Huntington's disease, and use as anticonvulsants.

IT 186495-99-8, NPS 1506

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(cyclic prolylglycine composition and therapeutic uses, and use with other agents)

RN 186495-99-8 CAPLUS

CN Benzene propanamine, 3-fluoro- $\gamma$ -(3-fluorophenyl)-N-methyl-, hydrochloride (9CI) (CA INDEX NAME)



● HCl

IT Bone morphogenetic proteins

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(2; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Glutamate receptors

RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(AMPA-binding, antagonists; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT CD antigens

RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(CD106, agents against; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT CD antigens

RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(CD11A, agents against; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT CD antigens

RL: BSU (Biological study, unclassified); BIOL (Biological study)

(CD18, agents against; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Proteins  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(FHF-1; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Proteins  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(FHF-2; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Proteins  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(FHF-3; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Proteins  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(FHF-4; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Nervous system, disease  
(Huntington's chorea; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Cell adhesion molecules  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(ICAM (intercellular adhesion mol.), agents against; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Integrins  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(LPAM-1 (lymphocyte Peyer's patch adhesion mol. 1), agents against; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Cell adhesion molecules  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(Leu-CAM (leukocytic cell adhesion mol.), agents against; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Cell adhesion molecules  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(MAdCAM-1 (mucosal addressin cell adhesion mol.-1), agents against; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Tumor necrosis factors  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(TNF- $\alpha$ ; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Cell adhesion molecules  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(VCAM-1 (vascular cell adhesion mol. 1), agents against; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Cerebrospinal fluid  
(artificial; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Brain  
(cerebellum, neurons; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Injury  
(cerebral; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Interferons  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(consensus $\chi$ ; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Surgery  
(coronary artery bypass, neurol. injury from; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Artery  
(coronary, bypass surgery, neurol. injury from; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Alzheimer's disease  
Anti-Alzheimer's agents  
Anti-inflammatory agents  
Antiparkinsonian agents  
Apoptosis  
Drug delivery systems  
Encephalomyelitis  
Glutamate antagonists  
Inflammation  
Multiple sclerosis  
Myelination  
Nerve  
Nerve, disease  
Nerve regeneration  
Nerve regeneration  
Nervous system agents  
Neuroglia  
Neuron  
Neurotoxicity  
Parkinson's disease  
Peptidomimetics  
(cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Ciliary neurotrophic factor  
Insulin-like growth factor-binding proteins  
Interleukins  
Leukemia inhibitory factor  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Nerve, disease  
(death; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Nervous system, disease  
(degeneration; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Neurotrophic factors  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(glial-derived; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Proteins  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(hst/Kfgk gene product; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Drug delivery systems  
(inhalants; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Drug delivery systems  
(injections, i.m.; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Drug delivery systems  
(injections, i.p.; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Drug delivery systems  
(injections, i.v.; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Drug delivery systems  
(injections, s.c.; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Drug delivery systems  
(injections; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Brain, disease

IT Nerve, disease  
(injury; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Proteins  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(int-2; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Brain, disease  
(leucodystrophy; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Glutamate receptors  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(metabotropic, mGluR2; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Glutamate receptors  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(metabotropic, mGluR3; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Antibodies and Immunoglobulins  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(monoclonal, MECA-367; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Axon  
(myelination; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Inflammation

IT Spinal cord, disease  
(myelitis, transverse; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Drug delivery systems  
(nasal; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Encephalitis  
(necrotizing hemorrhagic; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Central nervous system  
(neurogenesis; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Asphyxia  
(neurol. injury from; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Toxins  
RL: ADV (Adverse effect, including toxicity); BIOL (Biological study)  
(neurol. injury from; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Nervous system, disease  
(neuromyelitis optica; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Cell death  
(neuron; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Injury  
(neuronal; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Cytoprotective agents  
(neuroprotective; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Brain  
(nigrostriatum; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Inflammation

IT Nerve, disease  
(optic neuritis; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Drug delivery systems  
(oral; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Brain  
(pons, central pontine myelinolysis; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Brain, disease  
(progressive multifocal leukoencephalopathy; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Drug delivery systems  
(rectal; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Nervous system, disease  
(sclerosis, diffuse cerebral sclerosis of Schilder; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Proteins  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(somatotropin-binding; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Brain, disease  
(stroke, neurol. injury from; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Drug delivery systems  
(systemic; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Brain, disease  
(trauma; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Brain  
(white matter, damage; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Interferons  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
( $\gamma$ ; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Interferons  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
( $\alpha$ ; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Integrins  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
( $\alpha L$ , agents against; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Integrins  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
( $\alpha 4\beta 1$ , agents against; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Integrins  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
( $\alpha 4\beta 1$ , agents against; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Transforming growth factors  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
( $\beta 1$ ; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Interferons  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
( $\beta$ ; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT Integrins  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
( $\beta 7$ , agents against; cyclic prolylglycine composition and therapeutic uses, and use with other agents)

IT 492-27-3, Kynurenic acid 533-45-9, Clomethiazole 2578-57-6  
2578-57-6D, analogs and peptidomimetics 9002-72-6, Growth hormone  
9061-61-4, Nerve growth factor 14611-51-9, Selegiline 50913-82-1, ORG  
2766 67763-96-6, Insulin-like growth factor 1 67763-97-7, Insulin-like

growth factor 2 77086-21-6, Dizocilpine 77086-22-7, MK-801  
 80714-61-0, Semax 104987-11-3, FK506 106096-92-8, Acidic fibroblast  
 growth factor 106096-93-9, Basic fibroblast growth factor 106956-32-5,  
 Oncostatin M 109836-81-9, L-threo-1-Phenyl-2-decanoyleamino-3-morpholino-  
 1-propanol 114949-22-3, Activin 118876-58-7, NBQX 123584-45-2,  
 Fibroblast growth factor 4 130939-41-2, Fibroblast growth factor 6  
 130939-66-1, Neurotrophin 3 140698-57-3, Activity-dependent neurotrophic  
 factor 143375-33-1, Neurotrophin 4 148348-14-5, Fibroblast growth  
 factor 3 148348-15-6, Fibroblast growth factor 7 153436-22-7, GV  
 150526 161832-65-1, LY300164 161832-71-9, LY303070 171758-70-6,  
 Fibroblast growth factor 10 186495-99-8, NPS 1506 204719-95-9,  
 Fibroblast growth factor 16 524706-48-7, GV 1505260  
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
 (Biological study); USES (Uses)  
 (cyclic prolylglycine composition and therapeutic uses, and use with other  
 agents)

IT 56-86-0, L-Glutamic acid, biological studies  
 RL: ADV (Adverse effect, including toxicity); BSU (Biological study,  
 unclassified); BIOL (Biological study)  
 (glutamate toxicity; cyclic prolylglycine composition and therapeutic uses,  
 and use with other agents)

L3 ANSWER 8 OF 20 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2003:376586 CAPLUS  
 DOCUMENT NUMBER: 138:379245  
 TITLE: Cyclo(prolylglycine) and methods of use to treat  
 neural disorders  
 INVENTOR(S): Guan, Jian; Gluckman, Peter David; Sieg, Frank  
 PATENT ASSIGNEE(S): Neuronz Limited, N. Z.; Neuronz Biosciences, Inc.  
 SOURCE: PCT Int. Appl., 40 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 2  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2003039487	A2	20030515	WO 2002-US36235	20021112
WO 2003039487	A3	20040115		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GO, GW, ML, MR, NE, SN, TD, TG				

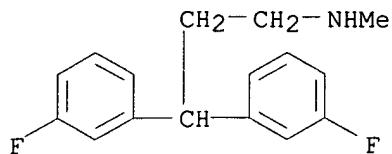
PRIORITY APPLN. INFO.: NZ 2001-515371 A 20011109  
                           NZ 2001-515432 A 20011113  
                           NZ 2001-515551 A 20011116

AB Embodiments of pharmaceutical compns. comprising cyclo(Pro-Gly) (cPG) and  
 methods for use in treating neural degeneration are provided. The cPG  
 substantially prevents toxic neural degeneration and cell death and  
 promotes neurite outgrowth in neurons, especially cerebellar neurons. The  
 neuroprotective and neuroregenerative effects of cPG are useful to treat  
 behavioral neurol. deficits involving motor control pathways.

IT 186495-99-8, NPS 1506  
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
 (Biological study); USES (Uses)  
 (cyclo(prolylglycine) for treatment of neural disorders, and use with  
 other agents)

RN 186495-99-8 CAPLUS

CN Benzenepropanamine, 3-fluoro-γ-(3-fluorophenyl)-N-methyl-,  
 hydrochloride (9CI) (CA INDEX NAME)



● HCl

- IT Bone morphogenetic proteins  
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (2; cyclo(prolylglycine) for treatment of neural disorders, and use with other agents)
- IT Glutamate receptors  
 RL: BSU (Biological study, unclassified); BIOL (Biological study)  
 (AMPA-binding, antagonists; cyclo(prolylglycine) for treatment of neural disorders, and use with other agents)
- IT Growth factors, animal  
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (Androgen-induced growth factor; cyclo(prolylglycine) for treatment of neural disorders, and use with other agents)
- IT CD antigens  
 RL: BSU (Biological study, unclassified); BIOL (Biological study)  
 (CD106; cyclo(prolylglycine) for treatment of neural disorders, and use with other agents)
- IT CD antigens  
 RL: BSU (Biological study, unclassified); BIOL (Biological study)  
 (CD11A; cyclo(prolylglycine) for treatment of neural disorders, and use with other agents)
- IT CD antigens  
 RL: BSU (Biological study, unclassified); BIOL (Biological study)  
 (CD18; cyclo(prolylglycine) for treatment of neural disorders, and use with other agents)
- IT Brain, disease  
 Nervous system, disease  
 (Devic's disease; cyclo(prolylglycine) for treatment of neural disorders)
- IT Proteins  
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (FHF-1; cyclo(prolylglycine) for treatment of neural disorders, and use with other agents)
- IT Proteins  
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (FHF-2; cyclo(prolylglycine) for treatment of neural disorders, and use with other agents)
- IT Proteins  
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (FHF-3; cyclo(prolylglycine) for treatment of neural disorders, and use with other agents)
- IT Proteins  
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (FHF-4; cyclo(prolylglycine) for treatment of neural disorders, and use with other agents)
- IT Growth factors, animal  
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (Glial-activating factor; cyclo(prolylglycine) for treatment of neural disorders, and use with other agents)
- IT Nervous system, disease  
 (Huntington's chorea; cyclo(prolylglycine) for treatment of neural

IT      disorders)

IT      Cell adhesion molecules

      RL: BSU (Biological study, unclassified); BIOL (Biological study)  
          (ICAM (intercellular adhesion mol.); cyclo(prolylglycine) for treatment  
          of neural disorders, and use with other agents)

IT      Integrins

      RL: BSU (Biological study, unclassified); BIOL (Biological study)  
          (LPAM-1 (lymphocyte Peyer's patch adhesion mol. 1);  
          cyclo(prolylglycine) for treatment of neural disorders, and use with  
          other agents)

IT      Cell adhesion molecules

      RL: BSU (Biological study, unclassified); BIOL (Biological study)  
          (Leu-CAM (leukocytic cell adhesion mol.); cyclo(prolylglycine) for  
          treatment of neural disorders, and use with other agents)

IT      Cell adhesion molecules

      RL: BSU (Biological study, unclassified); BIOL (Biological study)  
          (MAdCAM-1 (mucosal addressin cell adhesion mol.-1);  
          cyclo(prolylglycine) for treatment of neural disorders, and use with  
          other agents)

IT      Nervous system, disease

      (Machado-Joseph; cyclo(prolylglycine) for treatment of neural  
          disorders)

IT      Neuron

      (Purkinje cell, 5-fluorouracil- or cytosine arabinoside-induced loss of  
          Purkinje cells; cyclo(prolylglycine) for treatment of neural disorders)

IT      Tumor necrosis factors

      RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
          (Biological study); USES (Uses)  
          (TNF- $\alpha$ ; cyclo(prolylglycine) for treatment of neural disorders,  
          and use with other agents)

IT      Cell adhesion molecules

      RL: BSU (Biological study, unclassified); BIOL (Biological study)  
          (VCAM-1 (vascular cell adhesion mol. 1); cyclo(prolylglycine) for  
          treatment of neural disorders, and use with other agents)

IT      Encephalomyelitis

      (acute or chronic; cyclo(prolylglycine) for treatment of neural  
          disorders)

IT      Cerebrospinal fluid

      (artificial; cyclo(prolylglycine) for treatment of neural disorders)

IT      Neurotrophic factors

      RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
          (Biological study); USES (Uses)  
          (brain-derived; cyclo(prolylglycine) for treatment of neural disorders,  
          and use with other agents)

IT      Myelin

      RL: BSU (Biological study, unclassified); BIOL (Biological study)  
          (central pontine myelinolysis; cyclo(prolylglycine) for treatment of  
          neural disorders)

IT      Brain

      (cerebellum, cerebellar neuron; cyclo(prolylglycine) for treatment of  
          neural disorders)

IT      Brain

      (cerebellum, damage; cyclo(prolylglycine) for treatment of neural  
          disorders)

IT      Brain, disease

      (cerebellum, degeneration; cyclo(prolylglycine) for treatment of neural  
          disorders)

IT      Brain

      (cerebellum, hemorrhage; cyclo(prolylglycine) for treatment of neural  
          disorders)

IT      Brain

      (cerebellum, infarction; cyclo(prolylglycine) for treatment of neural  
          disorders)

IT      Injury

      (cerebral; cyclo(prolylglycine) for treatment of neural disorders)

IT      Surgery

      (coronary artery bypass; cyclo(prolylglycine) for treatment of neural  
          disorders)

IT      Artery

(coronary, bypass surgery; cyclo(prolylglycine) for treatment of neural disorders)

IT Brain  
(corpus striatum; cyclo(prolylglycine) for treatment of neural disorders)

IT Brain  
(cortex; cyclo(prolylglycine) for treatment of neural disorders)

IT Peptides, biological studies  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(cyclic; cyclo(prolylglycine) for treatment of neural disorders)

IT Alzheimer's disease  
Anti-Alzheimer's agents  
Anti-ischemic agents  
Antiparkinsonian agents  
Apoptosis  
Asphyxia  
Drug delivery systems  
Hypoxia  
Ischemia  
Motor skill disorders  
Necrosis  
Nerve  
Nervous system, disease  
Nervous system agents  
Neurotoxicity  
Parkinson's disease  
Wernicke-Korsakoff syndrome  
(cyclo(prolylglycine) for treatment of neural disorders)

IT Toxins  
RL: ADV (Adverse effect, including toxicity); BIOL (Biological study)  
(cyclo(prolylglycine) for treatment of neural disorders)

IT Anti-inflammatory agents  
Glutamate antagonists  
(cyclo(prolylglycine) for treatment of neural disorders, and use with other agents)

IT Ciliary neurotrophic factor  
Insulin-like growth factor-binding proteins  
Interleukins  
Leukemia inhibitory factor  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(cyclo(prolylglycine) for treatment of neural disorders, and use with other agents)

IT Nerve, disease  
(degeneration; cyclo(prolylglycine) for treatment of neural disorders)

IT Brain  
(dentate gyrus; cyclo(prolylglycine) for treatment of neural disorders)

IT Nervous system, disease  
(diffuse cerebral sclerosis of Schilder; cyclo(prolylglycine) for treatment of neural disorders)

IT Drugs  
(drug-induced cerebellar disorders; cyclo(prolylglycine) for treatment of neural disorders)

IT Endocrine system  
(endocrine cerebellar disorders; cyclo(prolylglycine) for treatment of neural disorders)

IT Neurotrophic factors  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(glial-derived; cyclo(prolylglycine) for treatment of neural disorders, and use with other agents)

IT Brain  
(hippocampus, CA4; cyclo(prolylglycine) for treatment of neural disorders)

IT Brain  
(hippocampus, sector CA1; cyclo(prolylglycine) for treatment of neural disorders)

IT Brain

(hippocampus, sector CA2; cyclo(prolylglycine) for treatment of neural disorders)

IT Brain  
(hippocampus, sector CA3; cyclo(prolylglycine) for treatment of neural disorders)

IT Proteins  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(hst/Kfgk gene product; cyclo(prolylglycine) for treatment of neural disorders, and use with other agents)

IT Drug delivery systems  
(inhalants; cyclo(prolylglycine) for treatment of neural disorders)

IT Drug delivery systems  
(injections, i.m.; cyclo(prolylglycine) for treatment of neural disorders)

IT Drug delivery systems  
(injections, i.p.; cyclo(prolylglycine) for treatment of neural disorders)

IT Drug delivery systems  
(injections, i.v.; cyclo(prolylglycine) for treatment of neural disorders)

IT Drug delivery systems  
(injections, s.c.; cyclo(prolylglycine) for treatment of neural disorders)

IT Drug delivery systems  
(injections; cyclo(prolylglycine) for treatment of neural disorders)

IT Brain, disease  
(injury; cyclo(prolylglycine) for treatment of neural disorders)

IT Proteins  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(int-2; cyclo(prolylglycine) for treatment of neural disorders, and use with other agents)

IT Metabolism  
(metabolic cerebellar disorders; cyclo(prolylglycine) for treatment of neural disorders)

IT Antibodies and Immunoglobulins  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(monoclonal, MECA-367; cyclo(prolylglycine) for treatment of neural disorders, and use with other agents)

IT Nervous system, disease  
(multiple system atrophy; cyclo(prolylglycine) for treatment of neural disorders)

IT Inflammation  
Spinal cord, disease  
(myelitis, transverse; cyclo(prolylglycine) for treatment of neural disorders)

IT Drug delivery systems  
(nasal; cyclo(prolylglycine) for treatment of neural disorders)

IT Encephalitis  
(necrotizing hemorrhagic; cyclo(prolylglycine) for treatment of neural disorders)

IT Nerve  
(neural fasciculation; cyclo(prolylglycine) for treatment of neural disorders)

IT Nervous system, disease  
(neuromyelitis optica; cyclo(prolylglycine) for treatment of neural disorders)

IT Cytoprotective agents  
(neuroprotective; cyclo(prolylglycine) for treatment of neural disorders)

IT Inflammation  
Nerve, disease  
(optic neuritis; cyclo(prolylglycine) for treatment of neural disorders)

IT Drug delivery systems  
(oral; cyclo(prolylglycine) for treatment of neural disorders)

IT Axon

IT      (outgrowth; cyclo(prolylglycine) for treatment of neural disorders)  
IT      Gelatins, biological studies  
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
      (pharmaceutical composition including; cyclo(prolylglycine) for treatment of  
      neural disorders)  
IT      Brain, disease  
      (progressive multifocal leukoencephalopathy; cyclo(prolylglycine) for  
      treatment of neural disorders)  
IT      Drug delivery systems  
      (rectal; cyclo(prolylglycine) for treatment of neural disorders)  
IT      Proteins  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
      (Biological study); USES (Uses)  
      (somatotropin-binding; cyclo(prolylglycine) for treatment of neural  
      disorders, and use with other agents)  
IT      Nervous system, disease  
      (spinocerebellar ataxia 1; cyclo(prolylglycine) for treatment of neural  
      disorders)  
IT      Nervous system, disease  
      (spinocerebellar ataxia 6; cyclo(prolylglycine) for treatment of neural  
      disorders)  
IT      Nervous system, disease  
      (spinocerebellar ataxia, 2; cyclo(prolylglycine) for treatment of  
      neural disorders)  
IT      Nervous system, disease  
      (spinocerebellar ataxia, 4; cyclo(prolylglycine) for treatment of  
      neural disorders)  
IT      Nervous system, disease  
      (spinocerebellar ataxia, 5; cyclo(prolylglycine) for treatment of  
      neural disorders)  
IT      Nervous system, disease  
      (spinocerebellar ataxia, 7; cyclo(prolylglycine) for treatment of  
      neural disorders)  
IT      Nervous system, disease  
      (spinocerebellar ataxia, dominantly or recessively inherited;  
      cyclo(prolylglycine) for treatment of neural disorders)  
IT      Nervous system, disease  
      (spinocerebellar ataxia, sporadic; cyclo(prolylglycine) for treatment  
      of neural disorders)  
IT      Brain, disease  
      (stroke; cyclo(prolylglycine) for treatment of neural disorders)  
IT      Brain, disease  
      (trauma; cyclo(prolylglycine) for treatment of neural disorders)  
IT      Interferons  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
      (Biological study); USES (Uses)  
      ( $\alpha$  and consensus; cyclo(prolylglycine) for treatment of neural  
      disorders, and use with other agents)  
IT      Interferons  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
      (Biological study); USES (Uses)  
      ( $\alpha$ ; cyclo(prolylglycine) for treatment of neural disorders, and  
      use with other agents)  
IT      Integrins  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
      ( $\alpha$ L; cyclo(prolylglycine) for treatment of neural disorders, and  
      use with other agents)  
IT      Integrins  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
      ( $\alpha$ 4; cyclo(prolylglycine) for treatment of neural disorders, and  
      use with other agents)  
IT      Integrins  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
      ( $\alpha$ 4 $\beta$ 1; cyclo(prolylglycine) for treatment of neural  
      disorders, and use with other agents)  
IT      Transforming growth factors  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
      (Biological study); USES (Uses)  
      ( $\beta$ 1-; cyclo(prolylglycine) for treatment of neural disorders, and

use with other agents)  
IT Interferons  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
( $\beta$ ; cyclo(prolylglycine) for treatment of neural disorders, and use with other agents)  
IT Integrins  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
( $\beta$ 7; cyclo(prolylglycine) for treatment of neural disorders, and use with other agents)  
IT 51-21-8, 5-Fluorouracil  
RL: ADV (Adverse effect, including toxicity); BIOL (Biological study)  
(5-fluorouracil-induced loss of Purkinje cells; cyclo(prolylglycine) for treatment of neural disorders)  
IT 64-17-5, Ethanol, biological studies  
RL: ADV (Adverse effect, including toxicity); BIOL (Biological study)  
(alc. cerebellar degeneration; cyclo(prolylglycine) for treatment of neural disorders)  
IT 56-40-6, Glycine, biological studies  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(binding; cyclo(prolylglycine) for treatment of neural disorders, and use with other agents)  
IT 3705-27-9  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(cyclo(prolylglycine) for treatment of neural disorders)  
IT 492-27-3, Kynurenic acid 533-45-9, Clomethiazole 9002-72-6, Growth hormone 9061-61-4, Nerve growth factor 14611-51-9, Selegiline 50913-82-1, ORG 2766 67763-96-6, IGF-1 67763-97-7, IGF-2 77086-22-7, MK-801 80714-61-0, Semax 104987-11-3, FK506 106096-92-8, Acidic fibroblast growth factor 106096-93-9, Basic fibroblast growth factor 106956-32-5, Oncostatin M 109836-81-9, L-threo-1-Phenyl-2-decanoylamino-3-morpholino-1-propanol 114949-22-3, Activin 118876-58-7, NBQX 123584-45-2, Fibroblast growth factor 4 130939-41-2, Fibroblast growth factor 6 130939-66-1, Neurotrophin 3 140698-57-3, Activity-dependent neurotrophic factor 143375-33-1, Neurotrophin 4 148348-14-5, Fibroblast growth factor 3 148348-15-6, Fibroblast growth factor 7 153436-22-7, GV 150526 161832-65-1, LY300164 161832-71-9, LY303070 171758-70-6, Fibroblast growth factor 10 186495-99-8, NPS 1506 204719-95-9, Fibroblast growth factor 16 524706-48-7, GV 1505260  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(cyclo(prolylglycine) for treatment of neural disorders, and use with other agents)  
IT 147-94-4, Cytosine arabinoside  
RL: ADV (Adverse effect, including toxicity); BIOL (Biological study)  
(cytosine arabinoside-induced loss of Purkinje cells; cyclo(prolylglycine) for treatment of neural disorders)  
IT 69-65-8, Mannitol 9004-54-0, Dextran, biological studies  
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(pharmaceutical composition including; cyclo(prolylglycine) for treatment of neural disorders)  
IT 57-41-0, Phenytoin  
RL: ADV (Adverse effect, including toxicity); BIOL (Biological study)  
(phenytoin-induced cerebellar atrophy; cyclo(prolylglycine) for treatment of neural disorders)

L3 ANSWER 9 OF 20 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2002:777693 CAPLUS  
DOCUMENT NUMBER: 137:299911  
TITLE: Neuroprotectant formulations  
INVENTOR(S): Hesson, David P.; Frazer, Glenn D.; Ross, Douglas  
PATENT ASSIGNEE(S): Neuron Therapeutics, Inc., USA  
SOURCE: PCT Int. Appl., 28 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002078670	A1	20021010	WO 2002-US5885	20020228
			W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG	
EP 1370240	A1	20031217	EP 2002-733809	20020228
	R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR		
US 2002193285	A1	20021219	US 2002-90441	20020304
PRIORITY APPLN. INFO.:			US 2001-331360P	P 20010302
			US 2001-798880	A 20010302
			WO 2002-US5885	W 20020228

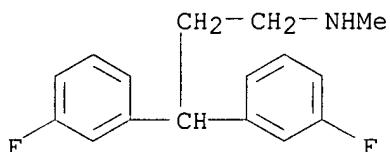
AB A method of treating an animal that has suffered damage to cerebrospinal tissue or that has an indication creating a risk of damage to cerebrospinal tissue, comprises injecting a physiol. acceptable cerebrospinal perfusion fluid into a first catheter into the cerebrospinal pathway. The cerebrospinal perfusion fluid has a neuroprotecting effective amount of a neuroprotectant, withdrawing fluid at a second catheter into the cerebrospinal pathway to create a flow and flow pathway between the first and second catheters and c. maintaining the flow for a period of time adapted to perfuse an affected tissue.

IT 186495-99-8, NPS 1506

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(neuroprotectant formulations)

RN 186495-99-8 CAPLUS

CN Benzenepropanamine, 3-fluoro- $\gamma$ -(3-fluorophenyl)-N-methyl-,  
hydrochloride (9CI) (CA INDEX NAME)



● HCl

IT Medical goods  
(catheters; neuroprotectant formulations)

IT Nervous system, disease  
(degeneration; neuroprotectant formulations)

IT Alzheimer's disease  
Anti-inflammatory agents  
Cerebrospinal fluid  
Drug delivery systems  
Human  
Multiple sclerosis  
Perfusion  
(neuroprotectant formulations)

IT Albumins, biological studies  
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(neuroprotectant formulations)

IT Cytoprotective agents  
(neuroprotective; neuroprotectant formulations)  
IT Anti-inflammatory agents  
(nonsteroidal; neuroprotectant formulations)  
IT Brain, disease

IT      (stroke; neuroprotectant formulations)  
IT      Injury  
IT      (trauma; neuroprotectant formulations)  
IT      169592-56-7, Caspase 3  
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
      (inhibitor; neuroprotectant formulations)  
IT      50-81-7, Ascorbic acid, biological studies 51-55-8, Atropine, biological  
      studies 533-45-9, Clomethiazole 987-78-0, Citicoline 2149-70-4,  
      Nitroarginine 2156-56-1, Ceresine 6735-59-7, Pralidoxime 19982-08-2,  
      Memantine 22059-21-8 22503-72-6 23052-81-5 23210-56-2, Ifenprodil  
      31409-32-2, MDL 27192 55985-32-5, Nicardipine 66085-59-4, Nimodipine  
      72784-43-1, ACPCM 72784-47-5, ACPCE 77086-21-6, Dizocilpine  
      79055-68-8 88191-84-8, MDL 28170 107452-89-1, Ziconotide  
      110347-85-8, Selfotel 111900-32-4 112924-45-5, Sinnabidiol  
      119431-25-3, Eliprotil 123931-04-4 125546-04-5 128073-45-0  
      128298-28-2, Remacemide 130931-65-6 137160-11-3, Cerestat  
      142852-51-5, TAK 147 144665-07-6, Lubeluzole 153504-81-5, Licostinel  
      158798-83-5, AK 275 160399-35-9, AK 295 168021-79-2, NXY 059  
      173952-44-8, SYM 2206 175615-45-9, LY 287041 185243-69-0, Etanercept  
**186495-99-8**, NPS 1506 223723-79-3, AEOL 10113 286475-30-7,  
      AEOL 10150 466685-97-2 466685-98-3  
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
      (neuroprotectant formulations)

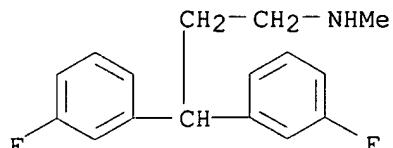
REFERENCE COUNT:      7      THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS  
                          RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3      ANSWER 10 OF 20     CAPLUS COPYRIGHT 2006 ACS on STN  
ACCESSION NUMBER:      2000:814825     CAPLUS  
DOCUMENT NUMBER:        135:55870  
TITLE:        NPS 1506 Attenuates Cognitive Dysfunction and  
                  Hippocampal Neuron Death Following Brain Trauma in the  
                  Rat  
AUTHOR(S):        Leoni, Matthew J.; Chen, Xiao-Han; Mueller, Alan L.;  
                  Cheney, Jessica; McIntosh, Tracy K.; Smith, Douglas H.  
CORPORATE SOURCE:       Department of Neurosurgery, University of  
                  Pennsylvania, Philadelphia, PA, 19104, USA  
SOURCE:        Experimental Neurology (2000), 166(2), 442-449  
                  CODEN: EXNEAC; ISSN: 0014-4886  
PUBLISHER:        Academic Press  
DOCUMENT TYPE:        Journal  
LANGUAGE:        English  
AB      Although several noncompetitive N-methyl-D-aspartate (NMDA) receptor  
      antagonists have been shown to be substantially efficacious in exptl.  
      models of brain trauma, side effects associated with this class of compds.  
      have impeded clin. application. Therefore, new noncompetitive NMDA  
      receptor antagonists have been developed, including NPS 1506, that appear  
      to be nontoxic but retain efficacy. In the present study, we evaluated  
      the efficacy of NPS 1506 in a model of parasagittal fluid percussion brain  
      trauma in the anesthetized rat. Administration of 1 mg/kg NPS 1506 at  
      both 10 min and 4 h posttrauma induced no changes in brain temperature, mean  
      arterial pressure, pulse, or arterial blood gasses. At 1 wk postinjury,  
      animals treated with the same dosing regimen of NPS 1506 demonstrated a  
      dramatic attenuation of memory dysfunction evaluated by a water maze task  
      and had greatly reduced neuron death in the CA3 subfield of the  
      hippocampus. However, NPS 1506 treatment did not significantly affect the  
      extent of cortical tissue loss following injury. Since memory dysfunction  
      and hippocampal damage are common and potentially related consequences of  
      brain trauma in humans, our results suggest that NPS 1506 treatment may  
      have clin. utility. (c) 2000 Academic Press.

IT      **186495-99-8**  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological  
study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES  
      (Uses)  
      (NPS 1506 attenuates cognitive dysfunction and hippocampal neuron death  
      following brain trauma in rats)

RN      186495-99-8     CAPLUS

CN      Benzenepropanamine, 3-fluoro- $\gamma$ -(3-fluorophenyl)-N-methyl-,  
      hydrochloride (9CI)    (CA INDEX NAME)



● HCl

IT Cognition enhancers  
 (NPS 1506 attenuates cognitive dysfunction and hippocampal neuron death following brain trauma in rats)

IT Memory, biological  
 (disorder; NPS 1506 attenuates cognitive dysfunction and hippocampal neuron death following brain trauma in rats)

IT Brain, disease  
 (hippocampus, injury; NPS 1506 attenuates cognitive dysfunction and hippocampal neuron death following brain trauma in rats)

IT Cytoprotective agents  
 (neuroprotectants; NPS 1506 attenuates cognitive dysfunction and hippocampal neuron death following brain trauma in rats)

IT Brain, disease  
 (trauma; NPS 1506 attenuates cognitive dysfunction and hippocampal neuron death following brain trauma in rats)

IT 186495-99-8  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (NPS 1506 attenuates cognitive dysfunction and hippocampal neuron death following brain trauma in rats)

REFERENCE COUNT: 57 THERE ARE 57 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 11 OF 20 CAPLUS COPYRIGHT 2006 ACS on STN  
 ACCESSION NUMBER: 2000:741905 CAPLUS  
 DOCUMENT NUMBER: 133:305610  
 TITLE: Treatment of neurological disorders with nitric oxide synthase inhibitors and excitatory amino receptor modulators  
 INVENTOR(S): O'Neill, Michael John  
 PATENT ASSIGNEE(S): Eli Lilly and Company Limited, UK  
 SOURCE: PCT Int. Appl., 22 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000061126	A2	20001019	WO 2000-GB1284	20000406
W:	AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			

PRIORITY APPLN. INFO.: GB 1999-8175 A 19990409

AB The present invention relates to a method of treating a neurol. disorder comprising administering to a patient an effective amount of a nitric oxide synthase inhibitor in combination with an effective amount of an excitatory amino receptor modulator. Combination of 2.5 mg/kg Mk-801, i.p., and 25

mg/kg ARL17477, i.p., had a synergistic degree of neuroprotection (78%) in cerebral ischemia induced in gerbils.

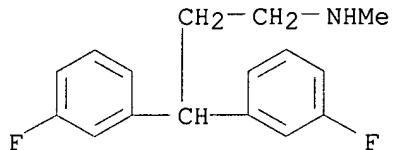
IT 186495-99-8, NPS 1506

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(treatment of neurol. disorders with nitric oxide synthase inhibitors and excitatory amino receptor modulators)

RN 186495-99-8 CAPLUS

CN Benzenepropanamine, 3-fluoro- $\gamma$ -(3-fluorophenyl)-N-methyl-, hydrochloride (9CI) (CA INDEX NAME)



● HCl

IT Glutamate receptors

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(AMPA-binding, antagonists; treatment of neurol. disorders with nitric oxide synthase inhibitors and excitatory amino receptor modulators)

IT Glutamate antagonists

(NMDA antagonists; treatment of neurol. disorders with nitric oxide synthase inhibitors and excitatory amino receptor modulators)

IT Nervous system

(disease; treatment of neurol. disorders with nitric oxide synthase inhibitors and excitatory amino receptor modulators)

IT Neurotransmitter antagonists

(excitatory amino acid; treatment of neurol. disorders with nitric oxide synthase inhibitors and excitatory amino receptor modulators)

IT Amino acids, biological studies

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(excitatory, agonists; treatment of neurol. disorders with nitric oxide synthase inhibitors and excitatory amino receptor modulators)

IT Brain, disease

(ischemia, focal; treatment of neurol. disorders with nitric oxide synthase inhibitors and excitatory amino receptor modulators)

IT Brain, disease

(ischemia; treatment of neurol. disorders with nitric oxide synthase inhibitors and excitatory amino receptor modulators)

IT Glutamate receptors

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(kainate-binding, antagonists; treatment of neurol. disorders with nitric oxide synthase inhibitors and excitatory amino receptor modulators)

IT Glutamate receptors

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(metabotropic, mGluR1, antagonists; treatment of neurol. disorders with nitric oxide synthase inhibitors and excitatory amino receptor modulators)

IT Glutamate receptors

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES

(Uses)  
(metabotropic, mGluR2, agonists; treatment of neurol. disorders with nitric oxide synthase inhibitors and excitatory amino receptor modulators)

IT Glutamate receptors  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES  
(Uses)  
(metabotropic, mGluR3, agonists; treatment of neurol. disorders with nitric oxide synthase inhibitors and excitatory amino receptor modulators)

IT Glutamate receptors  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES  
(Uses)  
(metabotropic, mGluR5, antagonists; treatment of neurol. disorders with nitric oxide synthase inhibitors and excitatory amino receptor modulators)

IT Cytoprotective agents  
(neuroprotectants; treatment of neurol. disorders with nitric oxide synthase inhibitors and excitatory amino receptor modulators)

IT 125978-95-2, Nitric oxide synthase  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(inhibitors; treatment of neurol. disorders with nitric oxide synthase inhibitors and excitatory amino receptor modulators)

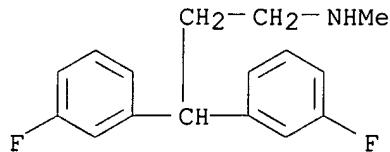
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25371-96-4, 1-(2-Trifluoromethylphenyl)imidazole 74209-34-0,  
3-Bromo-7-nitroindazole 77086-22-7, Mk-801 110347-85-8, Cgs 19755  
117414-74-1 118876-58-7, Nbqx 119431-25-3, Eliprodil 125546-04-5,  
Ly233053 137159-92-3, Aptiganel 137160-11-3, Cns1102 143343-70-8,  
Ly202157 143692-18-6, Ly300168 151056-97-2, 1701273 153436-38-5, GV  
150526A 153504-81-5, Acea1021 154164-30-4, Ym90k 154652-83-2,  
Ly293558 157971-06-7, GYK 152466 161832-65-1, Ly300164 168895-09-8,  
ARL17477 176199-48-7, Ly 354740 186495-99-8, NPS 1506  
191471-52-0, Ly379268 210245-80-0, Ym872 211566-75-5, Ly382884  
222529-89-7, LY 389795 301857-79-4, L-MIN 301857-80-7, Ramacemide  
301857-81-8, LY 377770  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES  
(Uses)  
(treatment of neurol. disorders with nitric oxide synthase inhibitors and excitatory amino receptor modulators)

L3 ANSWER 12 OF 20 CAPLUS COPYRIGHT 2006 ACS on STN  
ACCESSION NUMBER: 2000:740404 CAPLUS  
DOCUMENT NUMBER: 134:95398  
TITLE: NPS 1506, a moderate affinity uncompetitive NMDA receptor antagonist: preclinical summary and clinical experience  
AUTHOR(S): Mueller, A. L.; Artman, L. D.; Balandrin, M. F.; Brady, E.; Chien, Y.; DelMar, E. G.; Kierstead, A.; Marriott, T. B.; Moe, S. T.; Raszkiewicz, J. L.; Van Wagenen, B.; Wells, D.  
CORPORATE SOURCE: NPS Pharmaceuticals, Inc., Salt Lake City, UT, USA  
SOURCE: Amino Acids (2000), 19(1), 177-179  
CODEN: AACIE6; ISSN: 0939-4451  
PUBLISHER: Springer-Verlag Wien  
DOCUMENT TYPE: Journal  
LANGUAGE: English  
AB NPS Pharmaceuticals, Inc. (NPS) has synthesized a series of open-channel blockers with varying potencies at the NMDA receptor. NPS 1506 is a moderate affinity antagonist that inhibits NMDA/glycine-induced increases in cytosolic calcium in cultured rat cerebellar granule cells ( $IC_{50} = 476\text{nM}$ ) and displaces the binding of [ $^3\text{H}$ ]MK-801 to rat cortical membranes ( $IC_{50} = 664\text{nM}$ ).  
IT 186495-99-8, NPS 1506  
RL: ADV (Adverse effect, including toxicity); BPR (Biological process); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL

(Biological study); PROC (Process); USES (Uses)  
(NPS 1506, a NMDA receptor antagonist, preclin. summary and clin.  
experience)

RN 186495-99-8 CAPLUS

CN Benzenepropanamine, 3-fluoro- $\gamma$ -(3-fluorophenyl)-N-methyl-,  
hydrochloride (9CI) (CA INDEX NAME)



● HCl

IT Glutamate antagonists  
(NMDA antagonists; NPS 1506 a NMDA receptor antagonist, preclin.  
summary and clin. experience)

IT Cytoprotective agents  
(neuroprotectants; NPS 1506 a NMDA receptor antagonist, preclin.  
summary and clin. experience)

IT 186495-99-8, NPS 1506  
RL: ADV (Adverse effect, including toxicity); BPR (Biological process);  
BSU (Biological study, unclassified); THU (Therapeutic use); BIOL  
(Biological study); PROC (Process); USES (Uses)  
(NPS 1506, a NMDA receptor antagonist, preclin. summary and clin.  
experience)

L3 ANSWER 13 OF 20 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2000:381465 CAPLUS

DOCUMENT NUMBER: 133:30571

TITLE: Preparation of aralkylamines active at  
receptor-operated calcium channels as neuroprotectants

INVENTOR(S): Mueller, Alan L.; Balandrin, Manuel F.; Vanwagenen,  
Bradford C.; Delmar, Eric G.; Moe, Scott T.; Artman,  
Linda D.; Barmore, Robert M.

PATENT ASSIGNEE(S): NPS Pharmaceuticals, Inc., USA

SOURCE: U.S., 133 pp., Cont.-in-part of WO 9511663.

CODEN: USXXAM

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 6

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 6071970	A	20000606	US 1995-485038	19950607
CA 2182680	AA	19950817	CA 1994-2182680	19941026
WO 9521612	A2	19950817	WO 1994-US12293	19941026
WO 9521612	A3	19950921		
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CN 1148337	A	19970423	CN 1994-195074	19941026
CN 1088585	B	20020807		
ES 2156162	T3	20010616	ES 1994-932057	19941026
EP 1123922	A2	20010816	EP 2000-121960	19941026
EP 1123922	A3	20040102		
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PT 743853	T	20011031	PT 1994-932057	19941026
CA 2223978	AA	19961219	CA 1996-2223978	19960607

WO 9640097	A1	19961219	WO 1996-US10201	19960607
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ES, FI, GB, GE, HU, IS, JP, KE, KG, KP, KR, KZ, LK, LR, LS, LT,				
LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE,				
SG, SI				
RW: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
AU 9661125	A1	19961230	AU 1996-61125	19960607
AU 716122	B2	20000217		
EP 831799	A1	19980401	EP 1996-918477	19960607
EP 831799	B1	20030502		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,				
IE, FI				
CN 1192679	A	19980909	CN 1996-196042	19960607
JP 11506469	T2	19990608	JP 1996-502238	19960607
BR 9609019	A	19990706	BR 1996-9019	19960607
NZ 310344	A	20010330	NZ 1996-310344	19960607
AT 238782	E	20030515	AT 1996-918477	19960607
PL 185492	B1	20030530	PL 1996-323871	19960607
PT 831799	T	20030930	PT 1996-918477	19960607
ES 2197945	T3	20040116	ES 1996-918477	19960607
RU 2246300	C2	20050220	RU 1998-100454	19960607
US 6017965	A	20000125	US 1996-763480	19961211
HK 1008980	A1	20031107	HK 1998-109748	19980806
US 6211245	B1	20010403	US 1998-186341	19981104
US 6051610	A	20000418	US 1999-252433	19990218
AU 770292	B2	20040219	AU 2000-71810	20001124
US 2002004522	A1	20020110	US 2001-825373	20010402
US 6750244	B2	20040615		
JP 2004002437	A2	20040108	JP 2003-158350	20030603
US 2004171670	A1	20040902	US 2004-797355	20040309
PRIORITY APPLN. INFO.:			US 1993-14813	B2 19930208
			US 1994-194210	B2 19940208
			US 1994-288668	B2 19940809
			WO 1994-US12293	A2 19941026
			US 1994-288688	A2 19940811
			EP 1994-932057	A3 19941026
			JP 1995-521191	A3 19941026
			US 1995-485038	A 19950607
			US 1996-663013	A2 19960607
			WO 1996-US10201	W 19960607
			AU 1997-13525	A3 19961211
			US 1996-763480	A2 19961211
			US 1997-869154	B2 19970604
			US 1997-873011	A1 19970611
			US 1998-186341	A1 19981104
			US 2001-825373	A1 20010402

OTHER SOURCE(S): MARPAT 133:30571  
 AB Title compds., e.g., RCHR<sub>4</sub>CR<sub>1</sub>R<sub>5</sub>CR<sub>2</sub>R<sub>6</sub>R<sub>7</sub> [R = (un)substituted Ph; R<sub>1</sub>, R<sub>5</sub> = H, OH, (hydroxy)alkyl, alkoxy, acyloxy; R<sub>2</sub>, R<sub>6</sub> = H or hydroxyalkyl; R<sub>1</sub>R<sub>2</sub> = (CH<sub>2</sub>)<sub>n</sub> or (CH<sub>2</sub>)<sub>n</sub>NR<sub>3</sub>; R<sub>3</sub> = H, alkyl, CH<sub>2</sub>CH<sub>2</sub>OH; R<sub>4</sub> = (cyclo)alkyl, or (un)substituted Ph; R<sub>7</sub> = N(R<sub>3</sub>)<sub>2</sub>; R<sub>7</sub> = H when R<sub>1</sub>R<sub>2</sub> = (CH<sub>2</sub>)<sub>n</sub>NR<sub>3</sub>; n = 1-6] were prepared. Thus, (4-FC<sub>6</sub>H<sub>4</sub>)<sub>2</sub>CO was condensed with (EtO)<sub>2</sub>P(O)CH<sub>2</sub>CN and the product converted in 2 reduction steps to (4-FC<sub>6</sub>H<sub>4</sub>)<sub>2</sub>CHCH<sub>2</sub>CH<sub>2</sub>NH<sub>2</sub>. Data for biol. activity of title compds. were given.

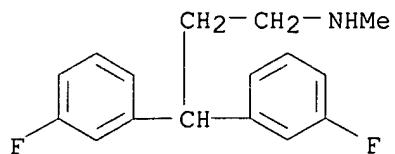
IT 186495-49-8P 186495-56-7P 186495-99-8P

273409-53-3P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (preparation of aralkylamines active at receptor-operated calcium channels as neuroprotectants)

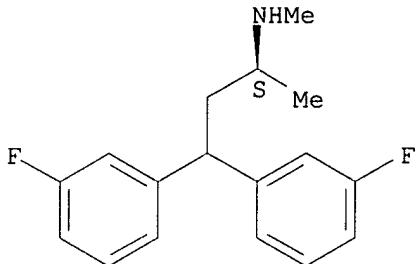
RN 186495-49-8 CAPLUS

CN Benzenepropanamine, 3-fluoro-γ-(3-fluorophenyl)-N-methyl- (9CI) (CA INDEX NAME)

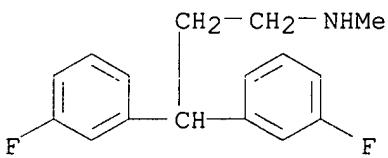


RN 186495-56-7 CAPLUS  
 CN Benzenepropanamine, 3-fluoro- $\gamma$ -(3-fluorophenyl)-N, $\alpha$ -dimethyl-,  
 ( $\alpha$ S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

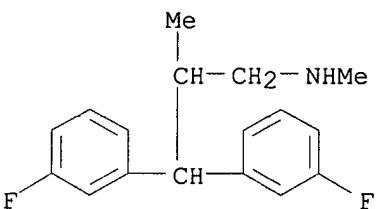


RN 186495-99-8 CAPLUS  
 CN Benzenepropanamine, 3-fluoro- $\gamma$ -(3-fluorophenyl)-N-methyl-,  
 hydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 273409-53-3 CAPLUS  
 CN Benzenepropanamine, 3-fluoro- $\gamma$ -(3-fluorophenyl)-N, $\beta$ -dimethyl-  
 (9CI) (CA INDEX NAME)



- IT Ionophores  
 (NMDA receptor complex; preparation of aralkylamines active at receptor-operated calcium channels as neuroprotectants)
- IT Glutamate receptors  
 RL: BSU (Biological study, unclassified); MSC (Miscellaneous); BIOL (Biological study)  
 (NMDA-binding, ionophore complex; preparation of aralkylamines active at receptor-operated calcium channels as neuroprotectants)
- IT Nervous system  
 (degeneration, treatment; preparation of aralkylamines active at receptor-operated calcium channels as neuroprotectants)

IT Cytoprotective agents  
 (neuroprotectants; preparation of aralkylamines active at receptor-operated calcium channels as neuroprotectants)

IT Calcium channel  
 RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
 (preparation of aralkylamines active at receptor-operated calcium channels as neuroprotectants)

IT 5586-73-2P 28075-29-8P 90531-05-8P 133805-32-0P 144451-98-9P  
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 217658-94-1P 217658-96-3P 217659-01-3P 217659-23-9P 217660-61-2P  
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**273409-53-3P**  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
 (preparation of aralkylamines active at receptor-operated calcium channels as neuroprotectants)

IT 62-23-7, p-Nitrobenzoic acid 85-41-6, Phthalimide 100-52-7,  
 Benzaldehyde, reactions 105-34-0, Methyl cyanoacetate 107-13-1,  
 2-Propenenitrile, reactions 109-76-2, 1,3-Diaminopropane 110-60-1,  
 1,4-Diaminobutane 135-02-4, o-Anisaldehyde 285-67-6, Cyclopentene  
 oxide 345-70-0, 3,3'-Difluorobenzophenone 443-73-2,  
 5-Fluoroindole-3-acetic acid 452-08-4, 2-Bromo-4-fluoroanisole  
 456-48-4, 3-Fluorobenzaldehyde 462-94-2, 1,5-Diaminopentane 529-20-4,  
 2-Methylbenzaldehyde 546-68-9 578-57-4, 2-Bromoanisole 587-04-2,  
 3-Chlorobenzaldehyde 932-31-0, 2-Methylphenylmagnesium bromide  
 1073-06-9, 1-Bromo-3-fluorobenzene 4587-33-1 5003-71-4 5460-29-7,  
 N-(3-Bromopropyl)phthalimide 7300-34-7, 4,9-Dioxa-1,12-dodecandiamine  
 17318-03-5, 3-Fluorophenylmagnesium bromide 17480-69-2,  
 (S)-N-Benzyl- $\alpha$ -methylbenzylamine 50715-13-4 65416-24-2, Benzyl  
 crotonate 77532-79-7, 5-Fluoro-2-methylbenzonitrile 122630-41-5  
 147624-13-3, 3-Fluoro-2-methylbenzaldehyde 168080-76-0,  
 3-Fluoro-2-methylbenzoyl chloride 263355-05-1, 3-Fluoro-2-  
 methylphenylmagnesium bromide  
 RL: RCT (Reactant); RACT (Reactant or reagent)  
 (preparation of aralkylamines active at receptor-operated calcium channels as neuroprotectants)

IT 455-67-4P 701-38-2P 4748-73-6P 14209-32-6P 35513-93-0P  
 38158-77-9P 51644-96-3P 75762-57-1P 83948-53-2P 98586-06-2P  
 101187-29-5P 114459-62-0P 122248-82-2P 122631-98-5P 122632-01-3P  
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**273409-62-4P**  
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)  
 (preparation of aralkylamines active at receptor-operated calcium channels as neuroprotectants)